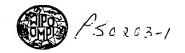
WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau



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(81) Designated States: JP, US, European patent (AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE).

Published

With international search report.

(54) Title: OXAZOLES FOR TREATING CYTOKINE MEDIATED DISEASES

(57) Abstract

This invention relates to the novel oxazole compounds of Formula (I) and novel pharmaceutical compositions comprising a compound of Formula (I) and a pharmaceutically acceptable diluent or carrier. This invention also relates to a method of inhibiting cytokines and the treatment of cytokine mediated diseases, in mammals, thereby by administration of an effective amount of a compound according to Formula (I).

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INTERNATIONAL SEARCH REPORT

International application No. PCT/US94/13067

Box I Observations where certain claims were found unsearchable (Continuation f item 1 of first sheet)	
This international report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:	
1. Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:	
2. Claims Nos.: because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:	
Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).	
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)	
This International Searching Authority found multiple inventions in this international application, as follows:	
Please See Extra Sheet.	
1. As all required additional search fees were timely paid by the applicant, this international search report covers all search claims.	able
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite paym of any additional fee.	nent
3. X As only some of the required additional search fees were timely paid by the applicant, this international search report coonly those claims for which fees were paid, specifically claims Nos.: 1-10(in part) of groups I, II and for asthma in claims 11-17	vers
4. No required additional search fees were timely paid by the applicant. Consequently, this international search report restricted to the invention first mentioned in the claims; it is covered by claims Nos.:	rt is
Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.	

INTERNATIONAL SEARCH REPORT

International application No. PCT/US94/13067

BOX II. OBSERVATIONS WHERE UNITY OF INVENTION WAS LACKING This ISA found multiple inventions as follows:

This application contains the following inventions or groups of inventions which are not so linked as to form a single inventive concept under PCT Rule 13.1. In order for all inventions to be examined, the appropriate additional examination fees must be paid.

I. Claims 1-10 wherein one of R1,R2 is pyridyl, the other R1,R2 and all other variables contain no additional heteroring.

II.Claims 1-10 wherein one of R1,R2 is pyrimidyl, the other R1,R2 and all other variables contain no additional heteroring.

III. Claims 1,4-8,10 wherein both R1,R2 contain pyridyl, other variables contain no additional hetero-ring.

IV. Claims 1,4-8, 10 wherein both R1, R2 contain pyrimidyl, other variables contain no additional hetero-ring.

V.Claims 1,4-8,10 wherein either or both R1,,R2 contain a hetero-ring other than pyridyl or pyrimidyl, each hetero ring or any combination of them is distinct. (Since the claimed language is indefinite, the exact meaning and scope can not be ascertained.) No additional hetero-ring anywhere in the formula.

VI. Claims 1,4-8, 10 wherein one or both R1, R2 contain hetero-ring other than pyridyl or pyrimidyl, each or combination thereof is distinct. (Other variables contain hetero-ring since the claimed language is indefinite. The exact meaning can not be ascertained.)

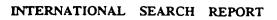
A method of use in claims 11-17 be considered together with any of the above group elected. The above groups lack unity of invention because the five-membered oxazole ring would not have been of sufficient similarity (common core structure) to allow a Markush grouping exhibiting unity. Note oxazole ring containing compounds exhibit a variety of uses such as sun screens.

INTERNATIONAL SEARCH REPORT

International application No. PCT/US94/13067

A. CLA	SSIFICATION OF SUBJECT MATTER					
IPC(6) :A61K 31/44,31/505; C07D 413/04						
	: 546/275; 544/331,333; 514/340,275,272,269,256		and IDG			
According to International Patent Classification (IPC) or to both national classification and IPC B. FIELDS SEARCHED						
2.						
Minimum d	ocumentation searched (classification system follower	d by classification s	ymbols)			
U.S. :	544/331,333 ;514/275,272,269,256,340; 546/275					
Documentat	tion searched other than minimum documentation to the	e extent that such do	cuments are included	in the fields searched		
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	data base consulted during the international search (n	ame of data base and	i, where practicable	, search terms used)		
APS,CA	S ON LINE					
C. DOC	CUMENTS CONSIDERED TO BE RELEVANT					
Category	Citation of document, with indication, where a	ppropriate of the re	evant nassages	Relevant to claim No.		
	, , , , , , , , , , , , , , , , , , , ,	ppropriate, or the re-	- Passages	Relevant to claim 140.		
X	Chemical Abstracts, volume 99, r	o.21,issued 2	1 November	1-17 -		
	1983, Sallmann, Trisubstituted	oxazo comp	ounds" see			
	abstract no. 175755w, WO 83/0			•		
	,			•		
X	Chemical Abstracts, volume 99, r	o.3. issued 18	3 July 1983	1-17		
	Sallmann ,"Diaryloxazole compo	nunds" soo (bstract no	1-17		
	22451u, S. African ZA 82/00,93	71103 366 6	abstract 110.			
	2240 Tu, 0. Amean 2A 02/00,93.	2 (12.2.02).				
x	Chamical Abstracts values 07 -	- 1 ! 10				
^ 1	Chemical Abstracts, volume 97, n Sallmann, "Trisubstituted	o. I, issued O		1-17		
-		imidazoie	derivatives:			
	pharmaceutical compositions cont	aining them ar	nd their use,"			
	see abstract no. 6298c, EP 45,08	11 (03.02.82)	•			
						
	er documents are listed in the continuation of Box C		ent family annex.			
· ·	ocial cotegories of cited documents:			mational filing date or priority tion but cited to understand the		
"A" doc	cument defining the general state of the art which is not considered be of particular relevance	principle or	theory underlying the inve			
.E. con	lier document published on or after the international filing date			e claimed invention cannot be red to involve an inventive step		
"L" doc	nument which may throw doubts on priority claim(s) or which is ed to establish the publication date of another citation or other		cument is taken alone			
apo	ocial reason (as specified)	"Y" document of	f particular relevance; the	claimed invention cannot be step when the document is		
"O. qoo	cument referring to an oral disclosure, use, exhibition or other	combined w		documents, such combination		
	nument published prior to the international filing date but later than priority date claimed	*&* document m	ember of the same patent	family .		
Date of the	Date of the actual completion of the international search Date of mailing of the international search Date of mailing of the international search					
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Commission	issioner of Patents and Trademarks					
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Form PCT/ISA/210 (second sheet)(July 1992)*



International application No. PCT/US94/13067

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C (Continua	tion). DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the releva	int passages	Relevant to claim N
X	Chemical Abstracts, volume 97, no.11, issued 13 Septe 1982, Sallmann, "Diarylmidazole compounds, pharmerodrugs containing them, and their use", see abstract no EP 46,451 (24.02.82).	ceutical	1-17
X	Chemical Abstracts, volume 100, no.1 issued 02 Januar Sallmann, "Trisubstituted oxazoles", see abstract no. 64 84,756(03.08.83).	y 1984, 96w, EP	1-17
X	Chemical Abstract, volume 83, no 15 issued 13 October Fitzi, "Imidazole derivatives" see abstract no. 131598n, 561,718 (15.05.75).	r, 1975, Swiss	1-17
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